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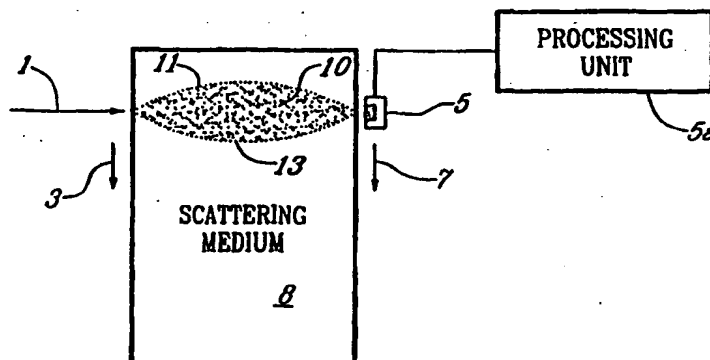|		
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(54) Title: DEPTH DISCRIMINATION OF HETOROGENEITIES IN TURBID MEDIA

(57) Abstract

In accordance with the method a turbid medium (8) is scanned by displacing an optical signal source and a corresponding optical detector (5) from one respective spatial location to another about the turbid medium, each spatial location being associated with corresponding input regions on opposite faces of the turbid medium (8). The optical signal source directs an optical signal (1) to each of said input regions. The optical detector (5) sends corresponding detector output signal data to an image processing means (5a) for generating a set of image data of the turbid medium (8). The improvement comprises generating in addition to a primary set of image data, secondary and/or tertiary image data which are derived by scanning the turbid medium (8) using input regions and output regions of different sizes. The various sets of image data obtained may be manipulated to highlight the differences or similarities between the images (e.g. to obtain a composite set of image data indicative of an anomaly(ies) in the body of the turbid medium (8).

DEPTH DISCRIMINATION OF HETOROGENEITIES IN TURBID MEDIA

The present invention relates to optical imaging of turbid media, e.g. breast tissue.

Optical mammography is likely to become a relevant technology for breast cancer detection.

5   Light scattering is the main problem that limits the potential of efficiently detecting small

lesions especially when located in the central region of the breast. Structures close to surface

are easily detected by optical techniques while more deep structures are more difficultly

detected. A technique to preferentially probe in the central region of the breast is thus of

interest.

10   The present text will refer by way of example only to breast tissue; however, other types of

turbid media (i.e. other types of (living) tissue) may also be imaged using the techniques

described herein.

Optical imaging of turbid media has been the subject of much research activity and has seen

an increase in interest since the early 1990's. This type of imaging is based on the fact that the

15   propagation of light in a turbid medium depends on the absorption and scattering properties

of the medium. Absorption results from energy level transitions of the constituent atoms and

molecules in the medium. The absorption property of the medium is quantified by its

absorption coefficient μ_a defined as the probability of a photon being absorbed per

infinitesimal pathlength. Scattering results from variations in the index of refraction of the

20   different structures present in the medium. In a highly diffusive medium, scattering is

quantified by the reduced scattering coefficient μ'_s, defined as the probability of a photon

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being isotropically scattered per infinitesimal pathlength. Characteristics such as intensity,

coherence and polarization of the incident light change as it is absorbed and scattered by the

medium resulting in diffuse transmittance of the light. In particular, scattering causes a

collimated laser beam to spread over a sizeable volume element. This complicates the

5   imaging of a turbid medium. Special imaging modalities must be implemented to offset the

detrimental light diffusion. For example, time-resolved methods use short light pulses from a

fast laser source to illuminate the medium. The emergent light is collected by a fast detector

capable of reproducing its time variation, which can provide further information about the

turbid medium. A simple data processing approach in this case is time-gating, by which only

10   the earliest part of the output light pulses is used to produce an image. This amounts to using

only the light with the straightest trajectory through the scattering medium, thus improving

spatial resolution (please see J. C. Hebden and R. A. Kruger, "Transillumination imaging

performance: Spatial resolution simulation studies", Med. Phys. 17, 41-47 (1990)) Reference

may also be made to the following documents for additional information with respect to

15   optical scanning techniques: S. B. Colak, D. G. Papaioannou, G. W. Hooft, M. B. Van der

Mark, H. Schomberg, J. C. J. Paasschens, J. B. M. Melissen, and N. A. A. J. Van Asten,

"Tomographic image reconstruction from optical projections in light-diffusing media", Appl.

Opt. 36, 180-213 (1997); M. S. Patterson, B. Chance, and B. C. Wilson, "Time-resolved

reflectance and transmittance for the noninvasive measurement of tissue optical properties",

20   Appl. Opt. 28, 2331-2336 (1989); D. Contini, F. Martelli, and G. Zaccanti, "Photon

migration through a turbid slab described by a model based on diffusion approximation. I.

Theory", Appl. Opt. 36, 4587-4599 (1997); M. Morin, S. Chatigny, A. Mailloux, Y.

Painchaud, and P. Beaudry, "Time-domain perturbation analysis of a scattering slab", in

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Optical Tomography and Spectroscopy of Tissue III, B. Chance, R. R. Alfano, and B. J.

Tromberg, eds., Proc. SPIE 3597, 67-78 (1999); Y. Painchaud, A. Mailloux, M. Morin, S.

Verreault and P. Beaudry, "Time-domain optical imaging: discrimination between scattering

and absorption", Appl. Opt. 38, 3686-3693 (1999); as well as U.S patent number 5,808,304).

5   The strong interest in optical imaging of scattering media stems from the need for biomedical

diagnostic techniques that are safe and non-invasive. The optical properties of biological

tissues are at the heart of optically based biomedical diagnostic techniques. As for the general

case of a turbid medium, the manner in which light propagates through tissue depends on its

absorption and scattering properties. Thus, if abnormal tissue can be said to differ from

10   normal in its absorption or scattering of light for some physiological or morphological reason,

it then becomes possible to optically differentiate between normal and abnormal conditions.

A specific application is optical mammography where tumors could be differentiated from

normal breast tissue on the basis of optical properties.

It would, for example, be advantageous to have a mechanism for the detection of an

15   anomaly(ies) in turbid medium such as for example a tumour in tissue such as breast tissue.

Accordingly, the present invention in one aspect provides in a method for scanning a turbid

medium for generating an image thereof for the detection of one or more anomalies contained

within the turbid medium, said turbid medium having a first face and an opposite second face,

wherein said turbid medium is scanned by displacing an optical signal source and a

20   corresponding optical detector from one respective spatial location to another about the

turbid medium, each spatial location being associated with a corresponding input region on

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said first face and a corresponding output region on the opposite second face, said optical

signal source directing an optical signal (e.g. laser beam) to each of said input regions, said

optical detector in response to optical signals detected from each of said output regions

sending corresponding detector output signal data to an image processing means for

5   generating a set of image data of the turbid medium, the improvement comprising

generating a primary set of image data derived by scanning said turbid medium using

for each input region a first input region of predetermined size and using for each output

region a first output region of predetermined size, and

generating a secondary set of image data derived by scanning said turbid medium

10   using for each input region said first input region of predetermined size and using for each

output

The present invention in another aspect provides in a method for scanning a turbid medium

for generating an image thereof for the detection of one or more anomalies contained within

the turbid medium, said turbid medium having a first face and an opposite second face,

15   wherein said turbid medium is scanned by displacing an optical signal source and a

corresponding optical detector from one respective spatial location to another about the

turbid medium, each spatial location being associated with a corresponding input region on

said first face and a corresponding output region on the opposite second face, said optical

signal source directing an optical signal (e.g. laser beam) to each of said input regions, said

20   optical detector in response to optical signals detected from each of said output regions

sending corresponding detector output signal data to an image processing means for

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generating a set of image data of the turbid medium, the improvement comprising

generating a primary set of image data derived by scanning said turbid medium using
for each input region a first input region of predetermined size and using for each output
region a first output region of predetermined size, and

5   generating a further set of image data derived by scanning said turbid medium using
for each input region a second input region of predetermined size greater than that of said first
input region and using for each output region said first output region of predetermined size.

The present invention in another aspect provides in a method for scanning a turbid medium
for generating an image thereof for the detection of one or more anomalies contained within

10   the turbid medium, said turbid medium having a first face and an opposite second face,
wherein said turbid medium is scanned by displacing an optical signal source and a
corresponding optical detector from one respective spatial location to another about the
turbid medium, each spatial location being associated with a corresponding input region on
said first face and a corresponding output region on the opposite second face, said optical

15   signal source directing an optical signal (e.g. laser beam) to each of said input regions, said
optical detector in response to optical signals detected from each of said output regions
sending corresponding detector output signal data to an image processing means for
generating a set of image data of the turbid medium, the improvement comprising

generating a primary set of image data I_A derived by scanning said turbid medium

20   using for each input region a first input region of predetermined size and using for each
output region a first output region of predetermined size,

generating a secondary set of image data I_B derived by scanning said turbid medium using for each input region said first input region of predetermined size and using for each output region a second output region of predetermined size greater than that of said first output region and

5　　　　generating a tertiary set of image data I_C derived by scanning said turbid medium using for each input region a second input region of predetermined size greater than that of said first input region and using for each output region said first output region of predetermined size.

In accordance with the present invention the various sets of image data may be generated in

10　　　any desired or necessary order.

In accordance with the present invention, the various sets of image data obtained as described herein may be manipulated in any manner (i.e. subjected to a data processing technique) so as to for example highlight the differences and/or similarities between the images (e.g.. to obtain a composite set of image data indicative of an anomaly(ies) (e.g. breast tumour) in the body

15　　　of the turbid medium); such manipulation or data processing treatment may be carried out by simple subtraction or by exploiting more sophisticated techniques such as for example K-L transforms. Thus, for example, the method(s) described herein may comprise (a) combining said primary set of image data and said secondary set of image data so as to generate a product set of image data wherein the similarities and/or differences between the

20　　　said primary set of image data and said secondary set of image data are highlighted.; (b) combining said primary set of image data and said further set of image data so as to generate

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a product set of image data wherein the similarities and/or differences between the said

primary set of image data and said further set of image data are highlighted.; (c) combining

said primary set of image data said secondary set of image data and said tertiary set of image

data so as to generate a product set of image data wherein the similarities and/or differences

5   between said primary set of image data, said secondary set of image data and said tertiary set

of image data are highlighted; etc.. Thus, more particularly, for example, a composite set of

image data may be generated by subtracting the secondary set of image data from the primary

set of image data; a composite set of image data may be generated by subtracting the further

set of image data from the primary set of image data; a composite set of image data may be

10   generated in accordance with the equation $I_B + I_C - I_A$..

In accordance with the present invention a system for performing optical imaging

measurements on a turbid medium in accordance with the method(s) as described herein may,

for example, comprise

any suitable (known) means for generating a collimated optical signal (e.g. laser)

15   directing means for directing said optical signal to an input area or region of

predetermined size on a face on one side of the turbid medium (e.g. using a suitable optical

fiber)

detection means for detecting a optical output signal from a second area or region of

predetermined size on the face of the turbid medium on the opposite side of said turbid

20   medium (e.g. photomuliplier tube, streak camera, etc..); the detection means may include

optical fibers for directing the optical signal to the component of the detection means which

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may convert the optical signal to an output (e.g. analogue or digital electrical) signal; and

means (e.g. computer) for processing said optical output signals (in any known manner) so as to obtain a set of image data (e.g. data which may be projected onto the screen of a computer monitor, be printed on a printer, be further processed, stored in computer
5   memory, etc...).

The system may also of course incorporate any suitable mechanism or means for manipulating the size of the input and output regions as referred to herein.

It is to be understood herein (i.e. for the purposes of the present invention) that the optical signal(s) may be of electromagnetic radiation which is not only in the visible but as well as in
10   the infra-red or the near infra-red part of the spectrum keeping in mind the purpose of the signal; the initial optical signal may for example be a laser beam.

The technique of the present invention may as mentioned above involve two or three (successive) scans of the turbid medium (e.g. breast tissue) so as to provide two or three projection images. These two or three images may be used to image structures according to
15   their depth inside the breast. The first image shows all the structures as in the usual optical techniques. A second image preferentially highlights the structures near the input surface. Another, third image preferentially highlights the structures near the output surface. By a combination of two or three of the images, it is possible to highlight the structures in the central region of the breast and attenuate the structures near the surfaces.

20   In the figures which illustrate example embodiments of the present invention:

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Figure 1 illustrates in schematic fashion a Scanning system for imaging through a turbid medium.

Figure 2 illustrates in schematic fashion a Scanning system for imaging through a turbid medium having a point-source/point-detector configuration

5   Figure 2A shows a graph for the configuration of Figure 2 wherein the qualitatively indicates the transmitted light versus transverse position for each configuration

Figure 3 illustrates in schematic fashion a Scanning system for imaging through a turbid medium having a point-source/large-area detector configuration

Figure 3A shows a graph for the configuration of Figure 3 wherein the qualitatively indicates

10   the transmitted light versus transverse position for each configuration

Figure 4 illustrates in schematic fashion a Scanning system for imaging through a turbid medium having a large-area source/point-detector configuration

Figure 4A shows a graph for the configuration of Figure 4 wherein the qualitatively indicates the transmitted light versus transverse position for each configuration;

15   Figure 5 illustrates a front view of a phantom;

Figure 6 illustrates a side view of the phantom of figure 5;

Figure 7 is a graphic illustration of Spatial profile combinations used to enhance longitudinal

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discrimination;

Figure 8 schematically illustrates an optical fiber arrangement for a point-source/point

detector configuration

Figure 9 schematically illustrates an optical fiber arrangement for a point-source/large-area

5   detector configuration

Figure 10 schematically illustrates an optical fiber arrangement for a large-area source/point-

detector configuration

Figure 11 is an illustration of the spatial profile combination for P_A-P_B of a liquid phantom

Figure 12 is an illustration of the spatial profile combination for P_A-P_C of a liquid phantom

10   Figure 13 is an illustration of the spatial profile combination for P_B + P_C -P_A of a liquid

phantom

Figure 14 is an illustration of the spatial profile for P_A of a liquid phantom

In biomedical optical imaging, two types of images can be generated: 3D reconstructed

images and 2D projection images. The former are produced using tomography which is

15   typically based on a multi-point geometry involving a large number of detectors (see S. B.

Colak, D. G. Papaioannou, G. W. Hooft, M. B. Van der Mark, H. Schomberg, J. C. J.

Paasschens, J. B. M. Melissen, and N. A. A. J. Van Asten, "Tomographic image

reconstruction from optical projections in light-diffusing media", Appl. Opt. 36, 180-213

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(1997)). Its advantage is that 3D images are generated, however, measurements and

reconstructions are potentially time-consuming.   2D projection images are based on the

detection of the light that enters the medium through a small area, that propagates through the

medium and that exits the medium over a small detection area facing the entrance area. The

5   image is generated by measuring the transmitted light in such a manner at a large number of

points as both the light beam incident on the medium and the detection device are scanned

synchronously along the surface of the medium. This imaging scheme is illustrated in Figure

1 wherein arrow 1 is indicative of a laser beam; wherein arrow 3 is indicative of the scanning

direction of the beam 1; wherein the optical detector means is indicated by reference numeral

10   5; wherein arrow 7 is indicative of the scanning direction of the optical detector means 5; and

wherein the scattering medium is designated by the reference numeral 8.   For the scanning

system schematically illustrated in figure 1 (known) means are provided (not shown) whereby

the laser beam 1 and the optical detector means 5 are respectively displaced in the direction of

the arrows 3 and 7 in synchronise fashion, i.e. such that the optical detector 5 is always

15   opposite the laser beam 1.   Although also not shown (in detail) in figure 1, the optical

detector 5 (as well as the means for generating the laser beam and the means for directing the

laser beam to the scan surface or face) may be coupled in any (known) manner to a suitable

(known) processing unit (e.g. computerised processing system) for ultimately generating or

deriving an image (i.e. a set of image data) from the set of signal data obtained from the

20   optical detector 5 during the scanning process.   Figure 1 does show in block diagram

fashion a processing unit 5a coupled to the optical detector 5 for processing the detector

output signal data into image data.

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The scanning technique illustrated by the system in figure 1 has the advantage that it is fast

and compatible with time-resolved measurements. However, information is limited to two

dimensions, the detected light giving information about a volume extending over the whole

line-of-sight joining the input point of the laser beam and the detector. This is illustrated in

5   Figure 1, where the gray region 10 within the dotted lines 11 and 13 represents the volume

through which detected photons have most likely propagated. The shape of this volume can

be understood by considering that all photons enter the scattering medium at the same point

and all detected photons leave it through a small area facing the detector. On the other hand,

scattering allows detected photons to wander away from the direct line-of-sight joining the

10   laser beam and the detector, this wandering being maximum at the half-distance between the

two. As illustrated in Figure 1, the scanning technique provides no longitudinal information,

since the probed volume extends over the whole thickness of the scattering medium.

Moreover, a small volume heterogeneity, with optical properties different from those of the

scattering medium, is more likely to be visible when located close to the input point of the

15   laser beam or the output point facing the detector. This is so because it is then probed by most

detected photons, whereas a smaller fraction of detected photons propagate through the same

heterogeneity standing in the middle of the scattering medium, where the probed volume is

the widest. For optical mammography this can be troublesome, as a tumor located deep

within breast tissue could be overshadowed by strongly absorbing structures such as veins

20   located near the surface.

The detection method in accordance with the present invention may provide for the basic

evaluation of heterogeneity depth in a scattering medium (e.g. turbid medium) to be used in

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conjunction with optical imaging based on light transmittance measurement and the

production of projection images. Depth assessment is necessary for a full 3-D localization of

the heterogeneity. It can also aid visualization of structures located in the central plane of a

turbid medium by identifying those structures located close to surfaces, which can then be

5   taken out of images. Finally, knowledge of the longitudinal position of the heterogeneity can

be used during data processing to improve the quantification of the optical contrast of the said

heterogeneity with regards to the surrounding scattering medium. Within the context of

mammography, this can help in tissue recognition.

10   The present invention which may be referred to by way of example as Dual Spatial

Integration (DSI) permits localization by separating the scattering medium into two or three

longitudinal zones: a zone close to the input surface where the light is injected, a central zone

and/or a zone close to the output surface where the light is detected. By dividing the medium

into two or three segments, structures close to surfaces can be extracted and set aside, and

15   structures located in the bulk enhanced.

Figure 2, 3 and 4 (as well as respective figures 2A, 3A and 4A) illustrate (in schematic

fashion) how detection surfaces and input beams of different sizes may be used to achieve a

certain degree of longitudinal localization of an inclusion within a scattering slab, e.g. by

20   appropriately modifying the input or output region sizes while maintaining the other

variables (i.e. essentially the same) during the generation of each of the sets of image data

during the scanning procedures.

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DSI may as mentioned above use three different source-detector configurations to achieve

this sectioning; example configurations are illustrated schematically in figures 2, 3 and 4; for

each of the shown configurations the same scattering medium is used, i.e. a scattering

medium containing three inclusions at different depths (i.e. the three black dots or circles

5   shown in these figures).

Specifically, the first configuration (i.e. configuration 1) shown in figure 2 comprises a point

source (i.e. small area input laser beam) 20 and a point detector (i.e. small area detector) 22

which are used as the base configuration.   The second configuration (i.e. configuration 2)

10   shown in figure 3 comprises a point source (i.e. small area input laser beam) 20 and a large

detector (i.e. large area detector) 24.   The third configuration (i.e. configuration 3) shown in

figure 4 comprises a large source (i.e. large area input laser beam) 26 and a point detector

(i.e. small area detector) 22 are used as the base configuration. For each of the configurations

the laser beam is generated in any known suitable fashion; the light leaving the scattering

15   medium is detected by any known suitable detector; the beam and detector (or alternatively,

the medium itself) are (is) displaced (i.e. so as to facilitate scanning of the medium)

longitudinally in synchronise fashion in any (known) suitable fashion across opposed faces

of the scattering medium; and images are generated from the detector signals by any suitable

known processing system (e.g. computer system). For each of figures 2, 3 and 4: the gray-

20   shaded rectangle is the turbid medium; the dark circles indicate the placement of the

inclusions or heterogeneities in the turbid medium; the dark line above each rectangle refers

to either a large area detector (long line) or a point detector (short line) - reference numbers

22 and 24; the arrows indicate the type of source used: single arrow for a point source and

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multiple arrows for a large area source, namely the elements specified by reference numbers
20 and 26; and the thin lines within the turbid medium delimitate qualitatively the volume
being probed by detected photons. The figures 2A, 3A and 4A show graphs wherein each
respectively qualitatively indicates the transmitted light versus transverse position for each
5   configuration.

In accordance with the present invention the detection method may proceed as discussed
hereinafter. The scattering medium is scanned with the scanning configuration shown in
figure 2 and an image (I_A) generated.   In the image (I_A) so obtained, structures close to both
10   the exit and entrance faces are highlighted relative to those within the bulk. . As the laser
beam and detector are moved transversely along the surface of the scattering medium, the
inclusions close to the input and output surfaces have a stronger impact on the detected light,
as discussed above.   This is represented qualitatively by the graph shown in figure 2A, which
depicts the detected light (signal) as a function of the position across the surface of the
15   scattering medium. A large area detector is then substituted for the point detector so as to
obtain the second configuration shown in figure 3. This changes the volume that is probed by
detected photons, as illustrated in Figure 3. The scattering medium is scanned with this
second configuration and an image (I_B) generated. In this image (I_B), structures closest to the
entrance face are highlighted, relative to those within the bulk or close to the exit face,
20   because they are probed by a larger fraction of the detected photons.   This is represented
qualitatively by the graph shown in figure 3A, which depicts the detected light (signal) as a
function of the position across the surface of the scattering medium. Thirdly, a large area
source is substituted for the point source and a point detector replaces the large area detector

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so as to obtain the third configuration. This changes again the shape of the volume being

probed by detected photons, as shown in Figure 4. The scattering medium is scanned with

this third configuration and an image (I_C) generated. In this image (I_C), structures closest to

the exit face are now highlighted relative to those within the bulk or close to the entrance

5   face. This is represented qualitatively by the graph shown in figure 4A, which depicts the

detected light (signal) as a function of the position across the surface of the scattering

medium.

If images I_B and I_C (i.e. their image or signal data) are added and image I_A (i.e. its image or

10   signal data) subtracted, structures located within the bulk of the scattering medium can be

more readily visualized than in images generated with the base configuration (I_A) alone.

It is to be understood herein that the expression "small area" is refers to a surface whose

diameter is relatively small compared to the scattering (turbid) medium thickness while the

15   expression "large area" is to be understood as referring to a surface whose diameter is on the

same order of magnitude as the scattering (turbid) medium thickness.

Application of DSI to the geometry illustrated in Figures 5 and 6 was evaluated numerically

and experimentally. As shown in figure 6 the scattering medium is a 5 cm-thick slab

20   containing three identical inclusions located at 5, 25 and 45 mm from the entrance face of the

slab. The inclusions are at a distance of 4 cm from one another in a direction parallel to the

face of the slab. The slab is 10 cm long (figure 5) with the inclusions spaced 4 cm apart from

the nearest neighbour inclusion.

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For the numerical simulations, the turbid medium was assumed to be purely scattering ($\mu_s =$ 1 mm^{-1}, $\mu_a = 0$ mm^{-1}). The inclusions were assumed to be point-like, i.e. to have a vanishingly small volume. They were also assumed to differ from the scattering medium only in their absorption coefficient, such that (volume)$_{inclusion}$ multiplied by $(\mu_a)_{inclusion} = 5$ mm^2.

5   The surface areas of both the large-area detector and source were set at 400 mm^2. The diffusion model (see M. S. Patterson, B. Chance, and B. C. Wilson, "Time-resolved reflectance and transmittance for the noninvasive measurement of tissue optical properties", Appl. Opt. 28, 2331-2336 (1989)), adapted to account for the refractive index mismatch between the turbid medium and the surrounding environment (see D. Contini, F. Martelli, and

10   G. Zaccanti, "Photon migration through a turbid slab described by a model based on diffusion approximation. I. Theory", Appl. Opt. 36, 4587-4599 (1997)), was used to describe light propagation through the simulated medium. The effect of the placement of an inclusion into the medium was modelled using a perturbation analysis (see M. Morin, S. Chatigny, A. Mailloux, Y. Painchaud, and P. Beaudry, "Time-domain perturbation analysis of a scattering

15   slab", in *Optical Tomography and Spectroscopy of Tissue III*, B. Chance, R. R. Alfano, and B. J. Tromberg, eds., Proc. SPIE 3597, 67-78 (1999)). The calculations assumed that infinitely short pulses of light were launched into the scattering medium. The transmitted light intensity calculation was limited to photons exiting the medium 2 ns after they entered the slab. The spatial intensity profiles P_A, P_B and P_C thus obtained with each detection

20   configuration were calculated for a transverse scan along a line joining the inclusions; P_A is profile for configuration 1, P_B is profile for configuration 2 and P_C is profile for configuration 3. These profiles were then normalized and combined to enhance depth discrimination, as discussed above. The results of these combinations are shown in Figure 7 after their

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normalization to unity.   Figure 7 is a graphic illustration of Spatial profile combinations used

to enhance longitudinal discrimination. The origin of the abscissa refers to the position of the

middle inclusion (see Figure 5: Front View). The value of x_{scan} refers to the displacement with

respect to this origin along the front surface of the medium (see Figure 3: Front View). P_x

5   refers to the spatial profiles calculated with scanning system configurations shown in figure 2,

3 and 4 (i.e. namely, configurations 1, 2 and 3).   In Figure 7 for the profile of P_A minus P_B is

illustrated by the graph line which contains the symbols ---○-- (designated by the reference

number 30); the profile of P_A minus P_C is illustrated by the graph line which contains the

symbols ---□-- (designated by the reference number 32); the profile of P_B plus P_C minus P_A

10   is illustrated by the graph line which does not contain any symbol (designated by the

reference number 34   ).

Referring to figure 7, as expected, when the profile for configuration 3 (P_C) is subtracted from

that of configuration 1 (P_A), only the signal from the inclusion near the input surface remains.

On the other hand, subtracting the profile for configuration 2 (P_B) from that of configuration 1

15   (P_A) nearly eliminates the signal change caused by the inclusion close to the input surface.

Figure 7 also shows the effect when the profiles for configurations 2 (P_B) and 3 (P_C) are

added together to get a resultant sum and the profile for configuration 1 (P_A) is subtracted

from the resultant sum.   These results clearly show how DSI permits to determine whether an

inclusion is located close to either surface of the scattering medium.

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The geometry in Figure 5 was investigated experimentally using a phantom consisting of

three inclusions placed into a 50 mm-thick cell containing a scattering solution of polystyrene

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microspheres. The inclusions were purely absorbing, 5 mm cylinders of black Delrin and

positioned according to the geometry illustrated in Figure 5. The scattering solution was non-

absorbing with $\mu = 1$ mm^{-1}. The laser light was carried close to the scattering medium using

an input optical fiber. Similarly, the light transmitted through the medium was collected with

5   an output optical fiber that carried the light to the detector. Using the divergence of light

emerging from a fiber, a large area source was simulated by simply moving the input fiber far

from the medium. Similarly, a large area detector was simulated by moving the output fiber

far from the medium. Moving the input fiber away from the phantom such that photons were

injected over a surface of 3.1 cm^2 functioned as the large area source. The large area detector

10   was likewise achieved by moving the output fiber away from the phantom such that photons

were collected over a surface of 3.8 cm^2.

The three example optical fiber arrangements are illustrated in Figure 8, 9 and 10. Figure 8

schematically illustrates an optical fiber arrangement for a point-source/point detector

configuration, i.e. the optical fibers are configured and spaced so as to provide a scanning

15   region or surface area and a detection region or surface area which have a small size and are

more or less the same size. Figure 9 schematically illustrates an optical fiber arrangement

for a point-source/large-area detector configuration i.e. the optical fibers 40 and 42 are

configured and spaced so as to provide a scanning region or surface area and a detection

region or surface area which are not the same size i.e. the detection area is larger than in

20   figure 8 - only the output fibers 42 are moved away from the phantom. Figure 10

schematically illustrates an optical fiber arrangement for a large-area source/point-detector

configuration i.e. the optical fibers are configured and spaced so as to provide a scanning

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region or surface area and a detection region or surface area which are not the same size i.e. the source area is larger than in figure 8 - only the output fibers 40 are moved away from the phantom. Although the figures show that the size of the input and output areas or regions may be varied by displacing the fibres 40 and 42 an other type of mechanism may of course

5 be used to effect the change in area. The phantom was scanned using each of these configurations using a near-infrared imaging system as described by Y. Painchaud, A. Mailloux, M. Morin, S. Verreault and P. Beaudry, "Time-domain optical imaging: discrimination between scattering and absorption", Appl. Opt. 38, 3686-3693 (1999)). The scan was done by moving synchronously both the input and output fibers 40 and 42

10 respectively along the surface of the medium in two dimensions in order to sequentially illuminate the entire surface. The detected signal was digitized and sent to a computer that was used to construct the images based of the series of measurements performed during a scan, each measurement corresponding to one pixel of the image.

Figure 11, 12 , 13 and 14 show the resultant projection images from scans of a liquid

15 phantom. The horizontal axis corresponds to the transversal placement of the inclusions. These images were normalized and combined as described earlier for the simulations. The point detector/point source configuration resulted in the image in Figure 14. Figures 11 and 12 show images demonstrating the success of the DSI method in extracting inclusions located near the surfaces. A comparison of the images in figures 13 and 14 shows that the inclusion

20 placed in the central plane of the phantom was highlighted relative to those near the surface using DSI.

20

As may be appreciated from the above, the Dual Spatial Integration technique (DSI) of the present invention is proposed as a tool to aid in the visualization of structures located in the central plane of a turbid medium. DSI represents an improvement with respect to the prior art in that one gains depth information content usually obtained with multi-point detection

5   geometry without its concomitant time costs. This is achieved by using different source-detector configurations to section the medium, namely as follows:

1. The scattering medium is scanned using a point source and point detector (i.e. configuration 1).

2. An image (I_A) is generated based on the detected light using configuration 1. As an

10   example, the detection of light can be done in a time-resolved manner in which only the first arrival photons are considered. The image can also be generated using indirect characteristics of the detected light as in the FIDM method described elsewhere[6.]

3. The scattering medium is scanned using a point source and a large area detector (configuration 2).

15   4. A second image (I_B) is generated using the same method as in 2 but with configuration 2.

5. The scattering medium is scanned using a large area source and a point detector (configuration 3).

6. A third image (I_C) is generated using the same method as in 2 but using the

20   configuration 3.

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●   ●

7. Image I_B is subtracted from image I_A to enhance structures located close to the output surface of the scattering medium.

8. Image I_C is subtracted from image I_A to enhance structures located close to the input surface of the scattering medium.

5   9. Image I_A is subtracted from the sum of images I_B and I_C to produce an image wherein the centrally-located structures are enhanced relative to those close to the surface.

Note that other combinations of images I_A, I_B and I_C or other data processing involving I_A, I_B and I_C may also result in obtaining relevant information.

In the summary given above, a point source refers to the injection of light in a medium over

10   a small area; a large area source refers to the injection of light in the medium over a large area; a point detector refers to the detection of light that emerges from the medium over a small area; a large area detector refers to the detection of light that emerges from the medium over a large area. Note again that "small area" refers to a surface whose diameter is small compared to the medium thickness while "large area" refers to a surface whose diameter is on

15   the same order of magnitude as the medium thickness. The appropriate input and output area sizes may be determined empirically for any given turbid medium to be subjected to analysis.

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We claim:

1.   In a method for scanning a turbid medium for generating an image thereof for the detection of one or more anomalies contained within the turbid medium, said turbid medium having a first face and an opposite second face, wherein said turbid medium

5   is scanned by displacing an optical signal source and a corresponding optical detector from one respective spatial location to another about the turbid medium, each spatial location being associated with a corresponding input region on said first face and a corresponding output region on the opposite second face, said optical signal source directing an optical signal to each of said input regions, said optical detector in

10   response to optical signals detected from each of said output regions sending corresponding detector output signal data to an image processing means for generating a set of image data of the turbid medium, the improvement comprising

generating a primary set of image data derived by scanning said turbid medium using for each input region a first input region of predetermined size and using for each output

15   region a first output region of predetermined size, and

generating a secondary set of image data derived by scanning said turbid medium using for each input region said first input region of predetermined size and using for each output region a second output region of predetermined size greater than that of said first output region.

20   2.   A method as defined in claim 1 comprising generating a composite set of image data by subtracting the secondary set of image data from the primary set of image data.

3.   In a method for scanning a turbid medium for generating an image thereof for the

detection of one or more anomalies contained within the turbid medium, said turbid

medium having a first face and an opposite second face, wherein said turbid medium

is scanned by displacing an optical signal source and a corresponding optical detector

5   from one respective spatial location to another about the turbid medium, each spatial

location being associated with a corresponding input region on said first face and a

corresponding output region on the opposite second face, said optical signal source

directing an optical signal to each of said input regions, said optical detector in

response to optical output signals detected from each of said output regions sending

10   corresponding detector output signal data to an image processing means for generating

a set of image data of the turbid medium, the improvement comprising

generating a primary set of image data derived by scanning said turbid medium using

for each input region a first input region of predetermined size and using for each output

region a first output region of predetermined size, and

15   generating a further set of image data derived by scanning said turbid medium using

for each input region a second input region of predetermined size greater than that of said first

input region and using for each output region said first output region of predetermined size.

·4   A method as defined in claim 3 comprising generating a composite set of image data

by subtracting the further set of image data from the primary set of image data.

20   5.   In a method for scanning a turbid medium for generating an image thereof for the

detection of one or more anomalies contained within the turbid medium, said turbid

24

medium having a first face and an opposite second face, wherein said turbid medium

is scanned by displacing an optical signal source and a corresponding optical detector

from one respective spatial location to another about the turbid medium, each spatial

location being associated with a corresponding input region on said first face and a

5   corresponding output region on the opposite second face, said optical signal source

directing an optical signal to each of said input regions, said optical detector in

response to optical output signals detected from each of said output regions sending

corresponding detector output signal data to an image processing means for generating

a set of image data of the turbid medium, the improvement comprising

10   generating a primary set of image data I_A derived by scanning said turbid medium

using for each input region a first input region of predetermined size and using for each

output region a first output region of predetermined size,

generating a secondary set of image data I_B derived by scanning said turbid medium

using for each input region said first input region of predetermined size and using for each

15   output region a second output region of predetermined size greater than that of said first

output region and

generating a tertiary set of image data I_C derived by scanning said turbid medium

using for each input region a second input region of predetermined size greater than that of

said first input region and using for each output region said first output region of
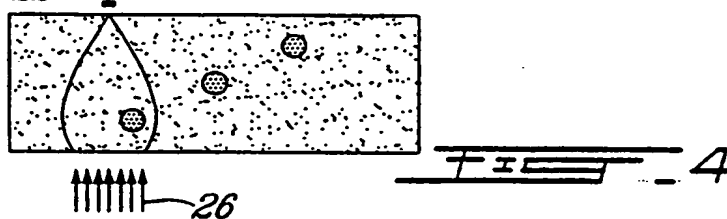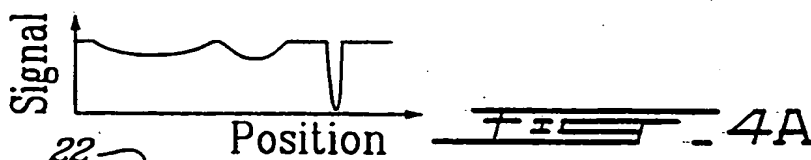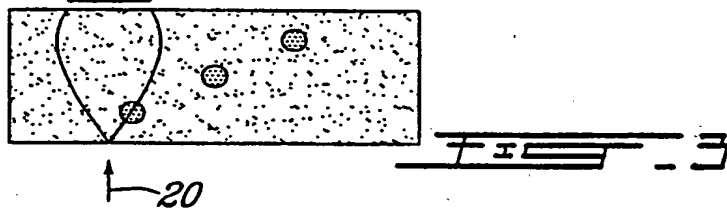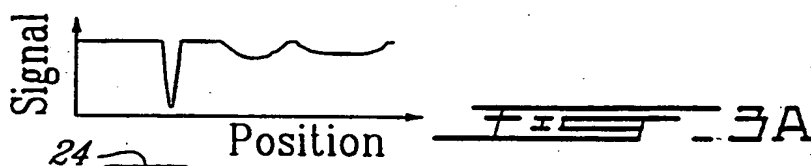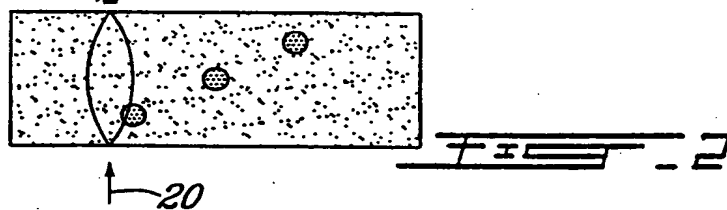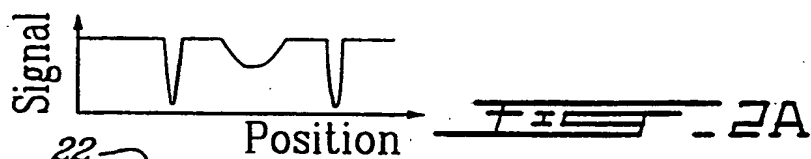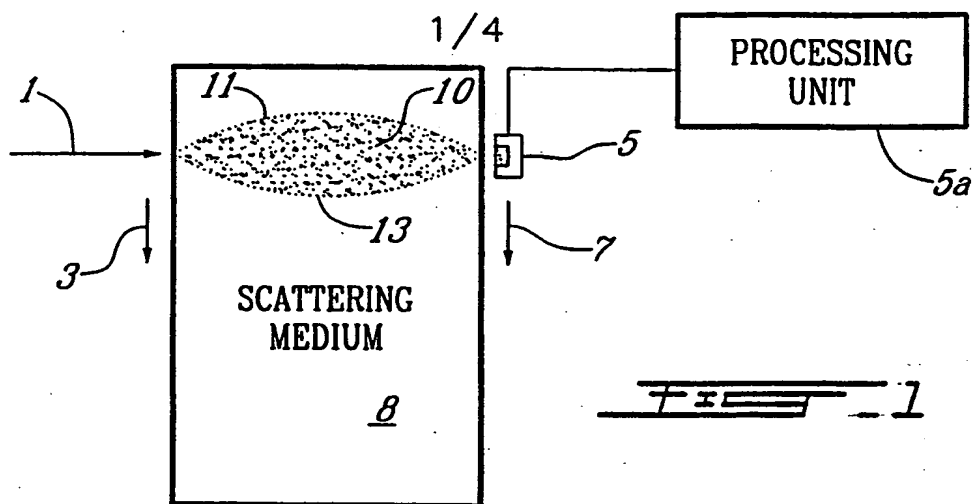
20   predetermined size.

6.   A method as defined in claim 5 comprising generating a composite set of image data

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in accordance with the equation $I_B + I_C - I_A$.

7.   A method as defined in claim 1 comprising combining said primary set of image data and said secondary set of image data so as to generate a product set of image data wherein the similarities and/or differences between the said primary set of image data and said secondary set of image data are highlighted.

8.   A method as defined in claim 3 comprising combining said primary set of image data and said further set of image data so as to generate a product set of image data wherein the similarities and/or differences between the said primary set of image data and said further set of image data are highlighted.

9.   A method as defined in claim 5 comprising combining said primary set of image data said secondary set of image data and said tertiary set of image data so as to generate a product set of image data wherein the similarities and/or differences between said primary set of image data, said secondary set of image data and said tertiary set of image data are highlighted.
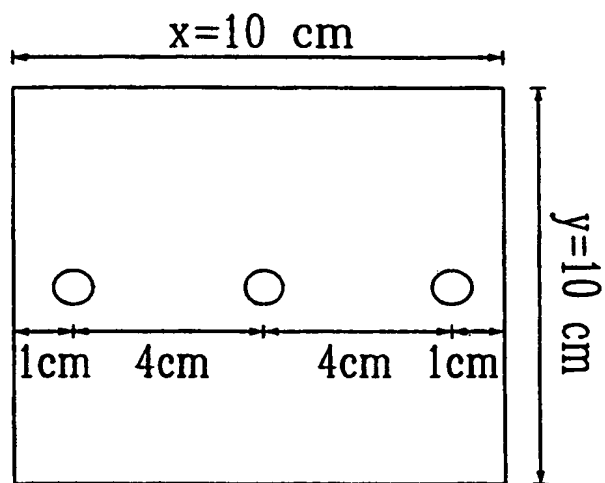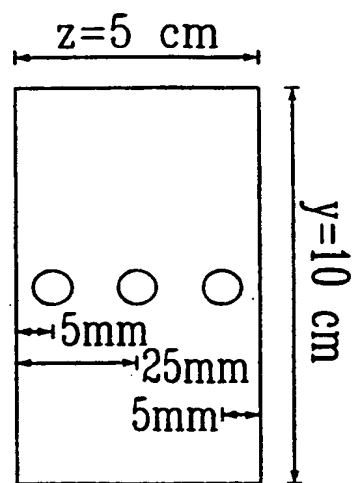
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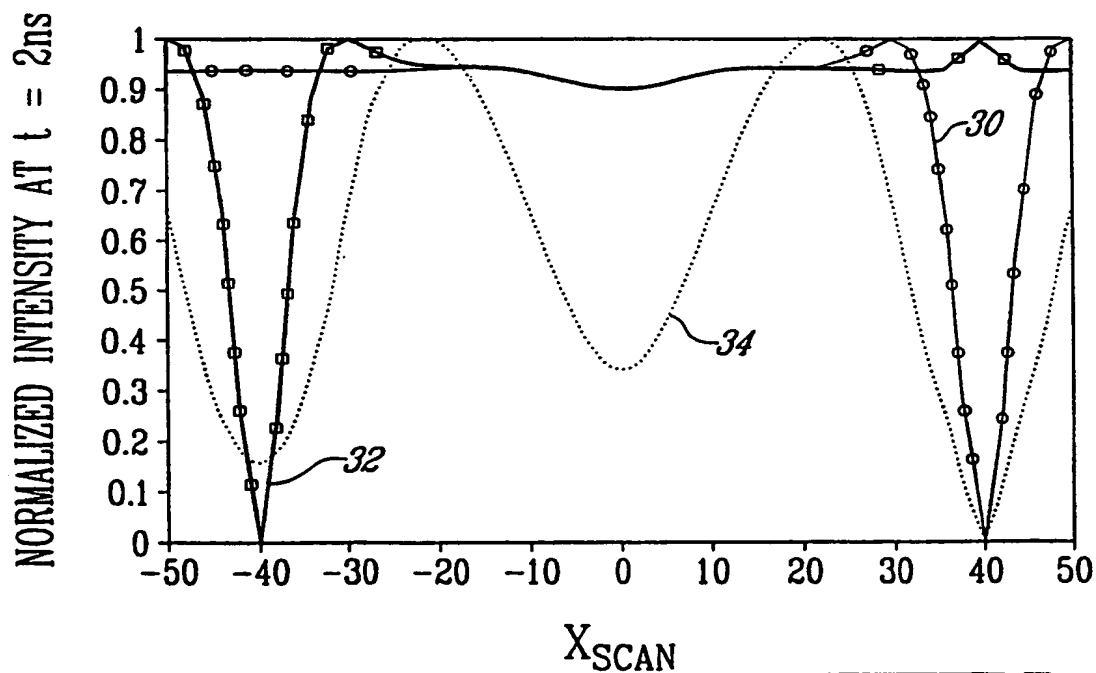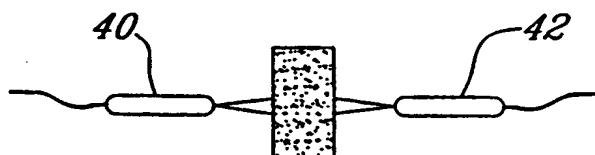PROCESSING UNIT

1

11   *10*

3

13

SCATTERING MEDIUM

8

5

5a

7

FIG. 1

Signal

Position

FIG. 2A

22

20

FIG. 2

Signal

Position

FIG. 3A

24

20

FIG. 3

Signal

Position

FIG. 4A

22

26

FIG. 4

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x=10 cm

y=10 cm

1cm   4cm   4cm   1cm

$F\llap{\rule[0.4ex]{1.5ex}{0.1ex}}I\rlap{\rule[0.4ex]{1.5ex}{0.1ex}}G.5$

z=5 cm

y=10 cm

5mm
25mm
5mm

$F\llap{\rule[0.4ex]{1.5ex}{0.1ex}}I\rlap{\rule[0.4ex]{1.5ex}{0.1ex}}G.6$

NORMALIZED INTENSITY AT t = 2ns

1
0.9
0.8
0.7
0.6
0.5
0.4
0.3
0.2
0.1
0

-50 -40 -30 -20 -10   0   10   20   30   40   50

X_{SCAN}

30

34

32

$F\llap{\rule[0.4ex]{1.5ex}{0.1ex}}I\rlap{\rule[0.4ex]{1.5ex}{0.1ex}}G.7$

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FIG. 8

FIG. 9

FIG. 10

FIG-11

FIG-12

FIG-13
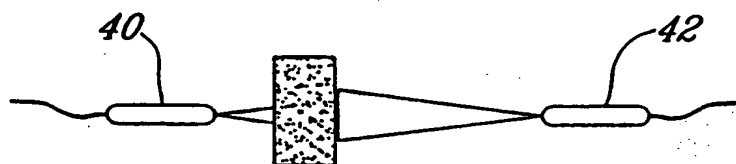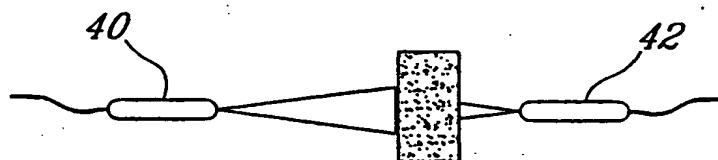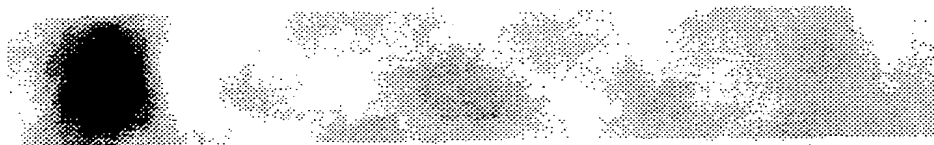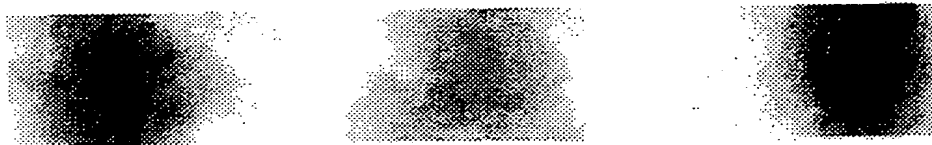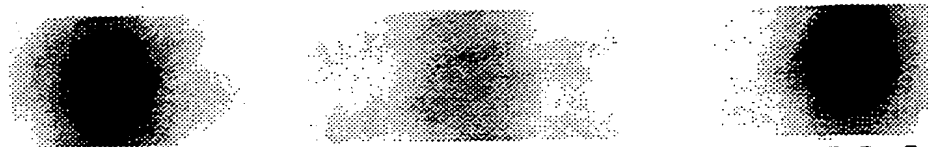
FIG-14

INTERNATIONAL SEARCH REPORT

INTERNATIONAL SEARCH REPORT

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7   A61B5/00   G01N21/47

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)
IPC 7 . A61B G01N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	DE 41 30 369 A (SIEMENS AG) 25 March 1993 (1993-03-25)	1
A	column 1, line 30 - line 53 column 4, line 24 -column 5, line 7	3,5
A	JARRY G ET AL: "LASER TOMOGRAPHY OF HETEROGENEOUS SCATTERING MEDIA USING SPATIAL AND TEMPORAL RESOLUTION" MEDICAL AND BIOLOGICAL ENGINEERING AND COMPUTING,GB,PETER PEREGRINUS LTD. STEVENAGE, vol. 31, no. 2, 1 March 1993 (1993-03-01), pages 157-164, XP000358964 ISSN: 0140-0118 sections "Devices and methods" and "Results"	1,3,5

-/--

[X] Further documents are listed in the continuation of box C.   [X] Patent family members are listed in annex.

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"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

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Date of the actual completion of the international search	Date of mailing of the international search report
3 May 2000	16/05/2000

Name and mailing address of the ISA	Authorized officer
European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo nl, Fax: (+31–70) 340–3016	Knüpling, M

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1

Internatonal Application No

PCT/CA 00/00052

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication,where appropriate, of the relevant passages	Relevant to claim No.
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EP 0108617	A	16-05-1984	AT	25815 T	15-03-1987
			DE	3370135 D	16-04-1987
			US	4600011 A	15-07-1986

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